

ABSTRACT

5 A method of treatment of disease by inhibition of cellular secretory
processes is provided. The method has particular application in the
treatment of diseases dependent upon the exocytotic activity of endocrine
cells, exocrine cells, inflammatory cells, cells of the immune system, cells
of the cardiovascular system, and bone cells. Agents and compositions
therefor, as well as methods for manufacturing these agents and
10 compositions, are provided. In a preferred embodiment a clostridial
neurotoxin, substantially devoid of holotoxin binding affinity for neuronal
cells of the presynaptic muscular junction, is associated with a targeting
moiety. The targeting moiety is selected such that the clostridial toxin
conjugate so formed may be directed to a non-neuronal target cell to which
15 the conjugate may bind. Following binding, a neurotoxin component of the
conjugate, which is capable of inhibition of cellular secretion, passes into
the cytosol of the target cell by cellular internalisation mechanisms.
Thereafter, inhibition of secretion from the target cell is effected.